

Anodal tDCS affects verbal episodic memory in initially low performers only

Habich A.¹, Klöppel S.^{1,2}, Peter J.^{1,2}

1 University Hospital of Old Age Psychiatry and Psychotherapy, University of Bern, Switzerland

2 Department of Psychiatry and Psychotherapy, Section of Gerontopsychiatry and Neuropsychology, Faculty of Medicine, University of Freiburg, Germany

Background

Episodic memory (i.e. connecting information with a specific context [1]) is highly relevant for the daily routine but undergoes decline during healthy aging and neurodegenerative disease. As the left dorsolateral prefrontal cortex (DLPFC) is active during both encoding and retrieval of episodic memories [2] it has become a frequent target in brain stimulation paradigms that boost neuronal excitability (i.e. transcranial direct current stimulation, tDCS [3,4]). However, there appears to be

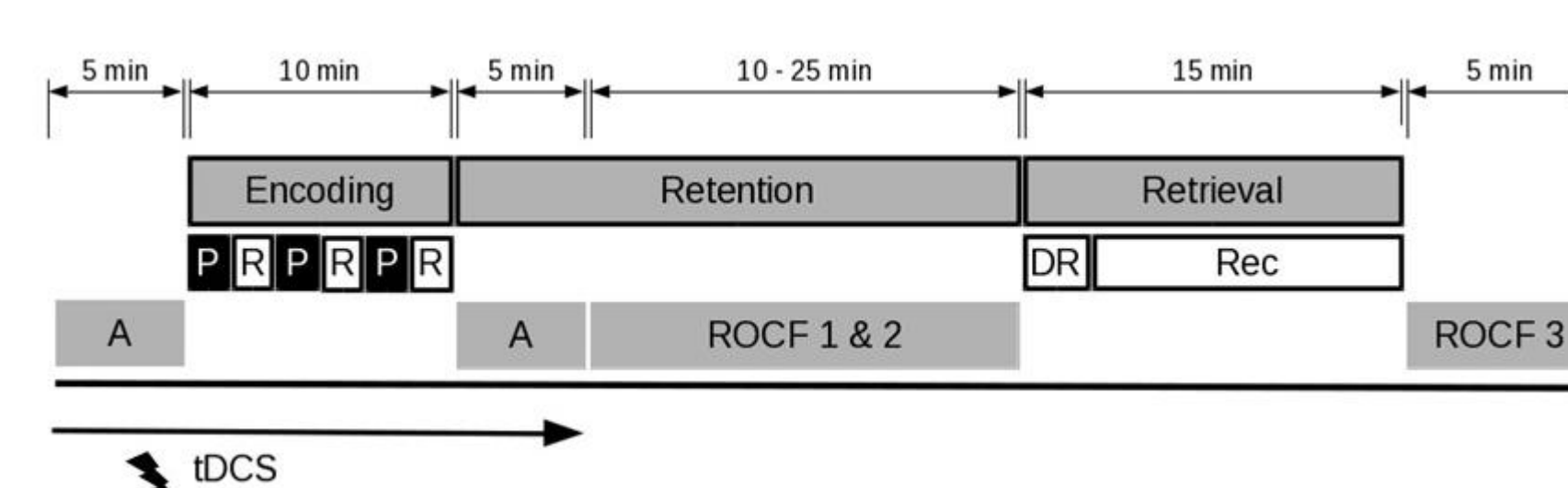
great variability in the tDCS response. Previous studies found that baseline performance influenced stimulation gains insofar as low performers were more likely to profit than their high performing peers. While this relationship has been mainly studied in a dichotomous manner, here we investigate if the modulating effect of tDCS on episodic memory is moderated by baseline performance even in a homogenous population of healthy young participants.

Methods

In this double-blind, sham-controlled study with parallel design, 43 healthy young adults (aged 24.8 ± 2.9 years, 22 females) received either 20 min of 1 mA anodal tDCS or sham stimulation to their left DLPFC during the encoding phase of a verbal episodic memory task (Fig. 1). During the encoding phase 40 nouns, extracted from the HVLt-R [5] were visually presented in three successive blocks, each of which was followed by an immediate oral retrieval. During the retention interval of approx. 20 min, participants completed an alertness task and furthermore copied and immediately retrieved the Rey-Osterrieth figure [6]. During the retrieval phase, participants first performed a delayed recall of the memorized words, and then completed a recognition task. Group differences induced by the stimulation were assessed by a 4 x 2 repeated measures ANOVA with retrieval (i.e. immediate and delayed recalls) as within-subject factor and stimulation as between-subjects factor. To investigate if initial performance moderated the benefit of tDCS on the increase in recalled words between first and last retrieval, we used

SPSS PROCESS macro [7] to perform a moderation analysis with stimulation (sham or real) as the focal predictor and midlist performance in the first retrieval as the moderator variable. Midlist performance in the first retrieval was chosen because it is uninfluenced by the serial position effect and thus better reflects baseline episodic memory ability [8].

Fig.1 Study procedure



Results

The overall fit of our moderation model reached significance ($F_{(3,39)} = 16.15$, $p < 0.0005$, $R^2 = 0.55$). Therein, the simple effects of initial midlist performance ($t_{(39)} = 5.2$, $p < 0.0005$), stimulation ($t_{(39)} = 2.74$, $p < 0.0009$) and their interaction ($t_{(39)} = -2.31$, $p = 0.03$) were significant. This indicates that initial low performers experienced the highest gain from stimulation on delayed recall (Fig. 3A, B).

The 4 x 2 repeated measures ANOVA showed only a significant main effect of retrieval ($F_{(3,38)} = 242.73$, $p < 0.0005$, $\eta^2 = 0.86$, Fig. 2) while no significant interaction between retrieval round and stimulation was found ($F_{(3,38)} = 2.18$, $p = 0.09$) on the group level.

Fig.2 Moderation effects

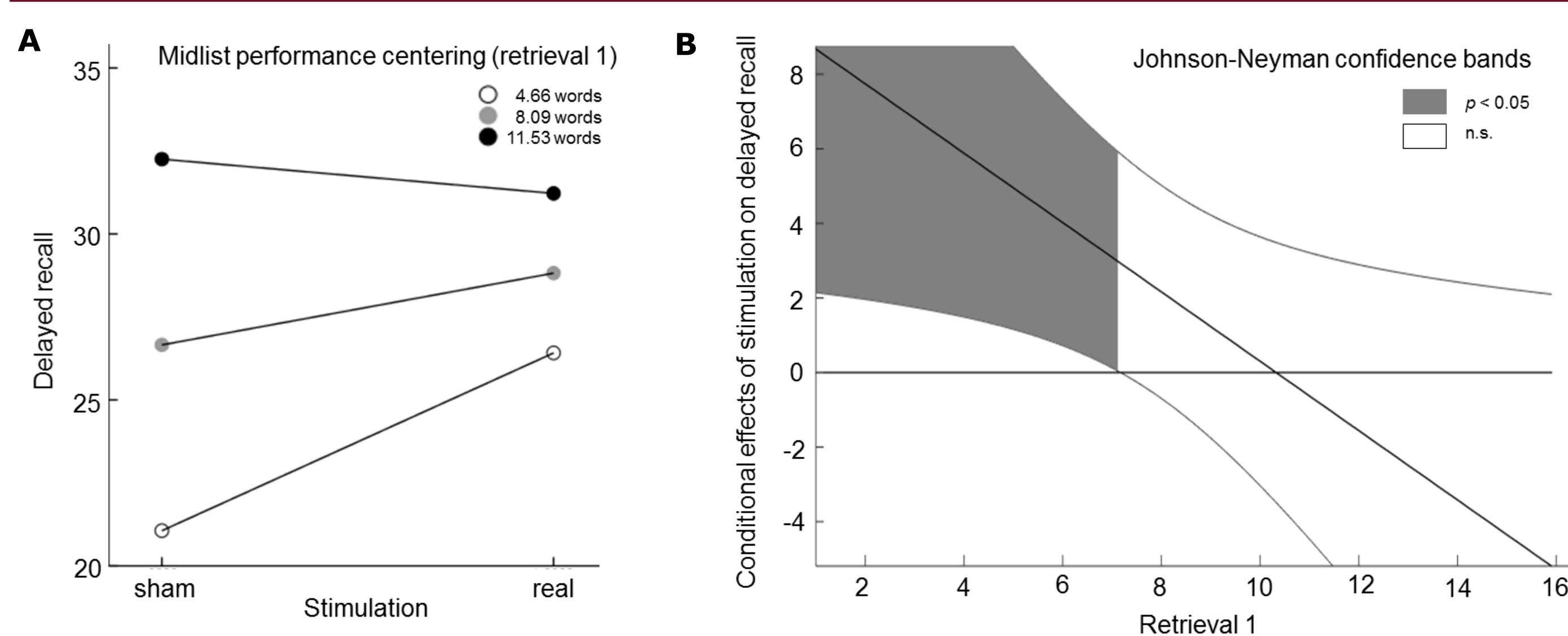
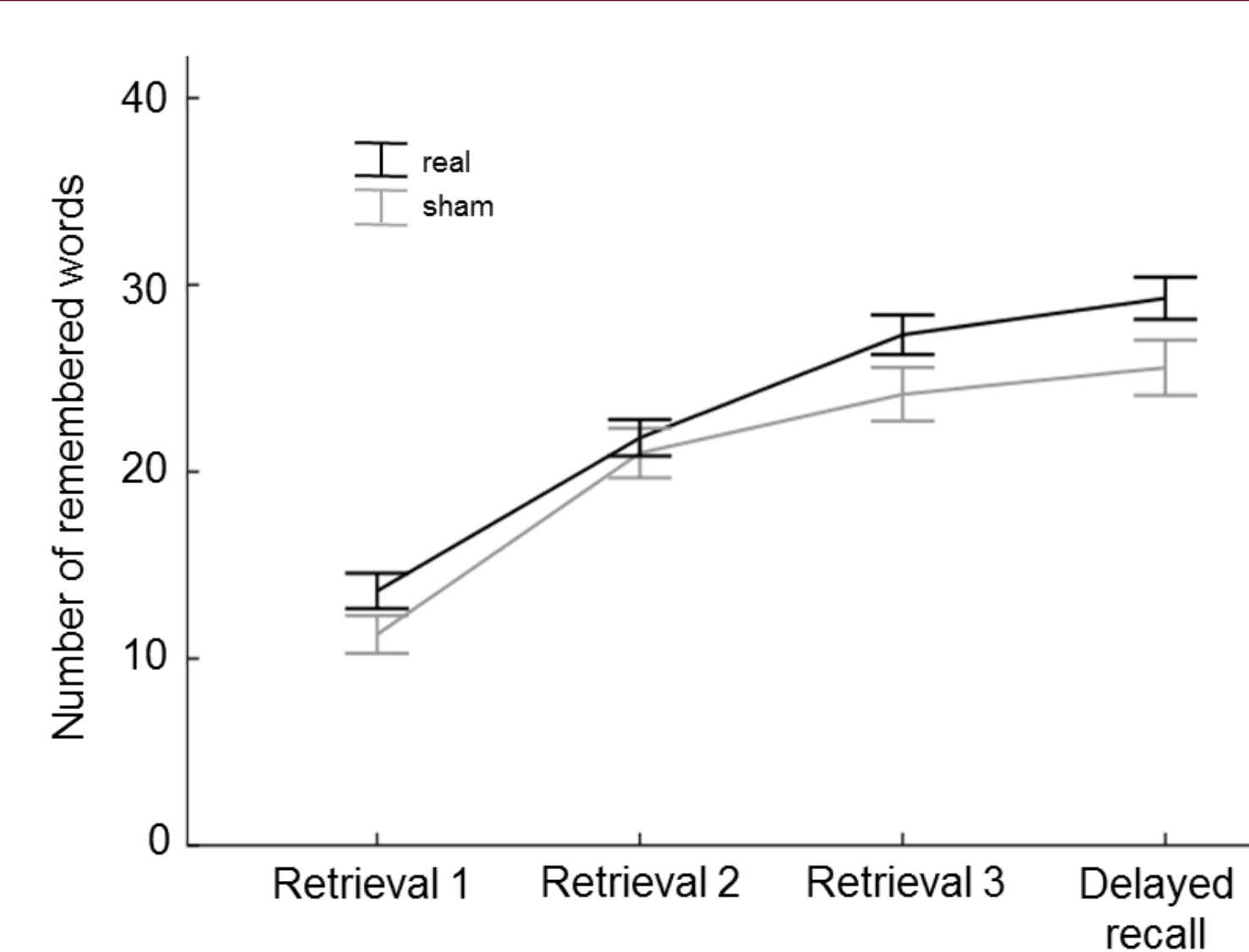


Fig. 3 Words retrieved during immediate and delayed recall



Conclusion

Notwithstanding that greater tDCS effect sizes have been attributed to both healthy older and neuropathologically afflicted populations [9], in the present study we showed that the favourable impact of tDCS on cognitive measures is not entirely restricted to specific populations. Individual baseline performance seems to be a more conclusive factor instead. Here, only initial low performers displayed tDCS-induced benefits in the delayed recall of a verbal episodic memory task. Similar inconsistencies regarding stimulation gains have been accumulating in the

literature. However, previous studies often divided their participants in two distinct groups, according to the level of expertise in the examined task, whereas the chosen continuous evaluation allows a more realistic representation of encountered abilities. Exposing the variability of brain stimulation outcomes provides valuable insights regarding basic research but also steers application-oriented research towards the customization of tDCS protocols, especially with regard to the identification of probable beneficiaries prior to the treatment.

- [1] Tulving E. Episodic Memory: From Mind to Brain. *Annu Rev Psychol.* 2002 Feb;53(1):1–25.
[2] Habib R et al. Hemispheric asymmetries of memory: the HERA model revisited. *Trends Cogn Sci.* 2003 Jun;7(6):241–5
[3] Manenti R et al. Enhancing verbal episodic memory in older and young subjects after non-invasive brain stimulation. *Front Aging Neurosci.* 2013 Sep 11;5:49
[4] Nikolai S et al. Focalised stimulation using high definition transcranial direct current stimulation (HD-tDCS) to investigate declarative verbal learning and memory functioning. *NeuroImage.* 2015 Aug;117:11–9.
[5] Benedict RHB et al. Hopkins Verbal Learning Test – Revised: Normative Data and Analysis of Inter-Form and Test-Retest Reliability. *Clin Neuropsychol. Neuropsychol Dev Cogn Sect D.* 1998 Feb 1;12(1):43–55
[6] Osterrieth PA. Le test de copie d'une figure complexe, contribution à l'étude de la perception et de la mémoire. *Archives de Psychologie.* 1944;40:206–356
[7] Hayes AF. Introduction to mediation, moderation, and conditional process analysis: a regression-based approach. New York: The Guilford Press; 2013. 507 p. (Methodology in the social sciences).
[8] Krueger LE. Influence of Cognitive Abilities and Age on Word Recall Performance Across Trials and List Segments. *Am J Psychol.* 2011;124(3):291–300
[9] Hsu, W et al. Effects of noninvasive brain stimulation on cognitive function in healthy aging and Alzheimer's disease: a systematic review and meta analysis. *Neurobiol. Aging* 2015 Aug; 36(8):2348–2359

Contact:

Annegret Habich
University Hospital of Old Age Psychiatry
University of Bern, Switzerland
E-Mail: annegret.habich@puk.unibe.ch

